## MARKED UP COPY OF ALL PENDING CLAIMS

- 1. (Amended herein) An indicator protein comprising:
  - a) a first binding moiety having a binding domain specific for a class of analytes that undergoes a reproducible allosteric change in conformation when said analytes are reversibly glucose bound;
  - d) a second moiety and third moiety that are covalently linked to either side of said first binding moiety in a manner that said second and third moieties undergo a change in relative position when said analyte molecule binds to said first binding moiety; and
  - e) said second and third moieties interact to produce a <u>fluorescent</u> change [in optical properties] when the relative positions of said second and third moieties change, wherein said <u>fluorescent</u> change can be monitored remotely by <u>external</u> optical means.

## 2. (Original) The protein of claim 1, wherein

- a) said first binding moiety is a protein that undergoes allosteric conformational changes when glucose reversibly binds;
- b) said second moiety is a fluorescent protein;
- c) said third moiety is a protein that has an absorption spectrum that overlaps the emission spectrum of said second moiety;
- f) the fluorescent energy transfer changes from said second moiety to said third moiety when glucose binds to said first binding moiety; and
- e) hybrid fusion joins said first, second and third moieties.

- 3. (Original) The protein of claim 2 wherein said third moiety is a fluorescent protein that can emit light when fluorescent energy transfers from said second moiety and said third moiety.
- 4. (Original) The protein of claim 2, wherein
  - b) said first binding moiety is a glucose binding protein from E. coli;
  - c) said second moiety is EBFP; and
  - d) said third moiety is hemoglobin.
- 5. (Amended herein) The protein of claim 2, wherein
  - a) said first binding moiety is a glucose binding protein from E. coli;
  - b) said second moiety is YFP; and
  - c) said third moiety [C] is GFP.
- 6. (Amended herein) The protein of claim 5 having the plasmid sequence shown in <u>SEQ</u>

  <u>ID NO: 1</u> [Figure 8].

- 7. (Amended herein) A biosensing system for glucose comprising:
  - d) a biosensor element consisting of a protein
    - having a first binding moiety, which is a glucose binding protein from E.
       coli, having a binding domain specific for glucose that undergoes a
       reproducible allosteric change when glucose is reversibly bound;
    - ii. having a second moiety and third moiety that are covalently linked to either side of said first binding moiety in a manner such that they change in relative position when glucose binds to said first binding moiety and wherein said second moiety and said third moiety interact to produce a fluorescent change [in optical properties] when their relative positions change wherein said fluorescent [optical properties] change can be monitored remotely by external optical means; and
    - iii. that is immobilized to a solid surface or retained within a permeable capsule;
  - e) the placement of said biosensor element in <u>subcutaneous</u> contact with a fluid of interest so that said biosensor element can be illuminated and emitted light detected; and
  - f) an <u>external</u> optical system for illumination of said biosensor element and detection of emitted radiation.
- (Original) A biosensing system for glucose of claim 7 wherein said second moiety is EBFP and said third moiety is hemoglobin.

- (Original) A biosensing system for glucose of claim 7 wherein said second moiety is
   YFP and said third moiety is GFP.
- 10. (Deleted) A biosensing system for glucose of claim 8 wherein said contact with a fluid of interest is subcutaneous.
- 11. (Deleted) A bionsensing system for glucose of claim 9 wherein said contact with said fluid of interest is subcutaneous.
- 12. (Original) A biosensing system for glucose of claim 8 wherein said contact with a fluid of interest occurs through a bioreactor.
- 13. (Original) A biosensing agent for glucose of claim 9 wherein said contact with a fluid of interest occurs through a bioreactor.
- 14. (Original) A biosensing system of claim 7 further comprising an instrument to measure changes in the fluorescence properties of said second moiety and said third moiety.

- 15. (Amended herein) A method for noninvasively measuring glucose within cells wherein
  - a. plasmid coding for a protein having
    - i. a first binding moiety having a binding domain specific for a class of analytes that undergoes a reproducible allosteric change in conformation when said analytes are reversibly glucose bound;
    - ii. a second moiety and third moiety that are covalently linked to either side of said first binding moiety in a manner that said second and third moieties undergo a <u>fluorescent</u> change in relative position when said analyte molecule binds to said first binding moiety; and
    - iii. said second and third moieties undergo a <u>fluorescent</u> change in optical properties when the relative positions of said second and third moieties, wherein said change can be monitored remotely by <u>external</u> optical means [. is] when introduced into cells;
  - b. said protein is expressed in the cells; and
  - c. said <u>fluorescent</u> changes [in fluorescence properties] are measured optically by an <u>external</u> instrument having an optical system for illumination and detection of emitted radiation.
  - 16. (Original) A method for noninvasively measuring glucose within cells of claim15 wherein said second moiety is YFP and said third moiety is GFP.

17. (Original) A method for noninvasively measuring glucose within cells of claim 15 wherein said second moiety is EBFP and said third moiety is hemoglobin.